Amendments to the Claims:

- 1-27. (Canceled).
- 28. (currently amended) An isolated nucleic acid <u>encoding a polypeptide</u> having at least 80% nucleic acid sequence identity to:
- (a) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334);
- (b) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
- (c) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334);
- (d) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
 - (e) the nucleic acid sequence shown in Figure 193 (SEQ ID NO:333);
- [[(f)]] (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333shown in Figure 193 (SEQ ID NO:333); or
- [[(g)]] (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270, wherein the encoded polypeptide induces chondrocyte proliferation.
- 29. (currently amended) The isolated nucleic acid of Claim 28 encoding a polypeptide having at least 85% nucleic acid sequence identity to:
- (a) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334 shown in Figure 194 (SEQ ID NO:334);
- (b) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
- (c) a nucleic acid-sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334);

- (d) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
 - (e) the nucleic acid sequence shown in Figure 193 (SEQ ID NO:333);
- [[(f)]] (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333shown in Figure 193 (SEQ ID NO:333); or
- [[(g)]] (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270, wherein the encoded polypeptide induces chondrocyte proliferation.
- 30. (currently amended) The isolated nucleic acid of Claim 28 encoding a polypeptide having at least 90% nucleic acid sequence identity to:
- (a) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334);
- (b) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
- (c) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEO ID NO:334);
- (d) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
 - (e) the nucleic acid sequence shown in Figure 193 (SEQ ID NO:333);
- [[(f)]] (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333shown in Figure 193 (SEQ ID NO:333); or
- [[(g)]] (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the encoded polypeptide induces chondrocyte proliferation.

- 31. (currently amended) The isolated nucleic acid of Claim 28 encoding a polypeptide having at least 95% nucleic acid sequence identity to:
- (a) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334sh);
- (b) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
- (c) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334);
- (d) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
 - (e) the nucleic acid sequence shown in Figure 193 (SEQ ID NO:333);
- [[(f)]] (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333 shown in Figure 193 (SEQ ID NO:333); or
- [[(g)]] (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270, wherein the encoded polypeptide induces chondrocyte proliferation.
- 32. (currently amended) The isolated nucleic acid of Claim 28 encoding a polypeptide having at least 99% nucleic acid sequence identity to:
- (a) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334);
- (b) a nucleic acid sequence encoding the amino acid sequence of the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
- (c) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334);
- (d) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;

- (e) the nucleic acid sequence shown in Figure 193 (SEQ ID NO:333);
- [[(f)]] (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333 shown in Figure 193 (SEQ ID NO:333); or
- [[(g)]] (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,
 - 33. (currently amended) An isolated nucleic acid comprising:

wherein the encoded polypeptide induces chondrocyte proliferation.

- (a) a nucleic acid sequence encoding the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334);
- (b) a nucleic acid sequence encoding the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
- (c) a nucleic acid sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334);
- (d)— a nucleic acid-sequence encoding the extracellular domain of the polypeptide shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide;
 - [[(e)]] (c) the nucleic acid sequence shown in Figure 193 (SEQ ID NO:333);
- [[(f)]] (d) the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333shown in Figure 193 (SEQ ID NO:333); or
- [[(g)]] (e) the full-length coding sequence of the cDNA deposited under ATCC accession number 203270.
- 34. (currently amended) The isolated nucleic acid of Claim 33 comprising a nucleic acid sequence encoding the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334).

- 35. (currently amended) The isolated nucleic acid of Claim 33 comprising a nucleic acid sequence encoding the polypeptide of SEQ ID NO:334shown in Figure 194 (SEQ ID NO:334), lacking its associated signal peptide.
 - 36. (canceled)
 - 37. (canceled)
- 38. (currently amended) The isolated nucleic acid of Claim 33 comprising the nucleic acid sequence of SEQ ID NO:333shown in Figure 193 (SEQ ID NO:333).
- 39. (currently amended) The isolated nucleic acid of Claim 33 comprising the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333 shown in Figure 193 (SEQ ID NO:333).
- 40. (previously presented) The isolated nucleic acid of Claim 33 comprising the full-length coding sequence of the cDNA deposited under ATCC accession number 203270.
 - 41. (canceled)
 - 42. (canceled)
 - 43. (canceled)
- 44. (currently amended) A vector comprising the nucleic acid of Claim 28, 48, 53 or 58.
- 45. (previously presented) The vector of Claim 44, wherein said nucleic acid is operably linked to control sequences recognized by a host cell transformed with the vector.
 - 46. (previously presented) A host cell comprising the vector of Claim 44.

- 47. (previously presented) The host cell of Claim 46, wherein said cell is a CHO cell, an *E. coli* or a yeast cell.
- 48. (new) An isolated nucleic acid encoding a polypeptide having at least 80% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the encoded polypeptide induces chondrocyte re-differentiation.

- 49. (new) The isolated nucleic acid of Claim 48 encoding a polypeptide having at least 85% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the encoded polypeptide induces chondrocyte re-differentiation.

- 50. (new) The isolated nucleic acid of Claim 48 encoding a polypeptide having at least 90% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;

- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the encoded polypeptide induces chondrocyte re-differentiation.

- 51. (new) The isolated nucleic acid of Claim 48 encoding a polypeptide having at least 95% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the encoded polypeptide induces chondrocyte re-differentiation.

- 52. (new) The isolated nucleic acid of Claim 48 encoding a polypeptide having at least 99% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the encoded polypeptide induces chondrocyte re-differentiation.

- 53. (new) An isolated nucleic acid encoding a polypeptide having at least 80% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide stimulates the uptake of glucose or FFA (free fatty acid) by adipocyte cells.

- 54. (new) The isolated nucleic acid of Claim 53 encoding a polypeptide having at least 85% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide stimulates the uptake of glucose or FFA (free fatty acid) by adipocyte cells.

- 55. (new) The isolated nucleic acid of Claim 53 encoding a polypeptide having at least 90% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;

- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide stimulates the uptake of glucose or FFA (free fatty acid) by adipocyte cells.

- 56. (new) The isolated nucleic acid of Claim 53 encoding a polypeptide having at least 95% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334,
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide stimulates the uptake of glucose or FFA (free fatty acid) by adipocyte cells.

- 57. (new) The isolated nucleic acid of Claim 53 encoding a polypeptide having at least 99% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or

(d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide stimulates the uptake of glucose or FFA (free fatty acid) by adipocyte cells.

- 58. (new) An isolated nucleic acid encoding a polypeptide having at least 80% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide induces differentiation of pancreatic β -cell precursor cells.

- 59. (new) The isolated nucleic acid of Claim 58 encoding a polypeptide having at least 85% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide induces differentiation of pancreatic β -cell precursor cells.

60. (new) The isolated nucleic acid of Claim 58 encoding a polypeptide having at least 90% sequence identity to:

- (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide induces differentiation of pancreatic β -cell precursor cells.

- 61. (new) The isolated nucleic acid of Claim 58 encoding a polypeptide having at least 95% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or
- (d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide induces differentiation of pancreatic β -cell precursor cells.

- 62. (new) The isolated nucleic acid of Claim 58 encoding a polypeptide having at least 99% sequence identity to:
 - (a) the amino acid sequence of the polypeptide of SEQ ID NO:334;
- (b) the amino acid sequence of the polypeptide of SEQ ID NO:334, lacking its associated signal peptide;
- (c) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the nucleic acid sequence of SEQ ID NO:333; or

(d) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 203270,

wherein the polypeptide induces differentiation of pancreatic β -cell precursor cells.

Please charge any additional fees, including any fees for additional extension of time, or credit overpayment to Deposit Account No. <u>08-1641</u> (referencing Attorney's Docket No. <u>39780-2830 P1C62</u>).

Respectfully submitted,

Date: January 6, 2005

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SV 2092595 v1 1/6/05 11:02 AM (39780.2830)